(Sheet 1 of 28)

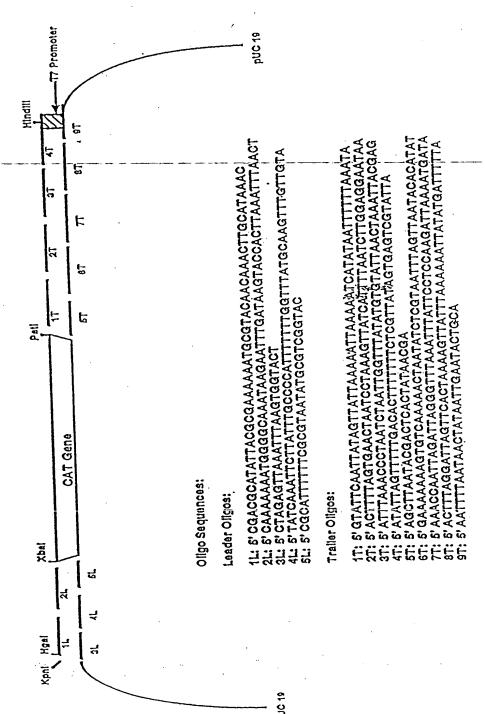
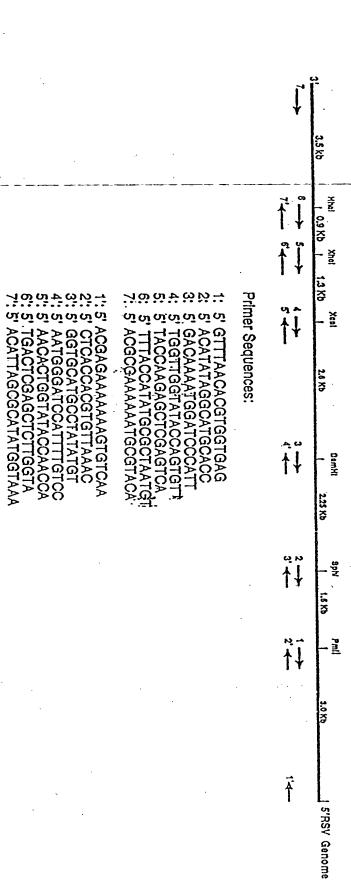
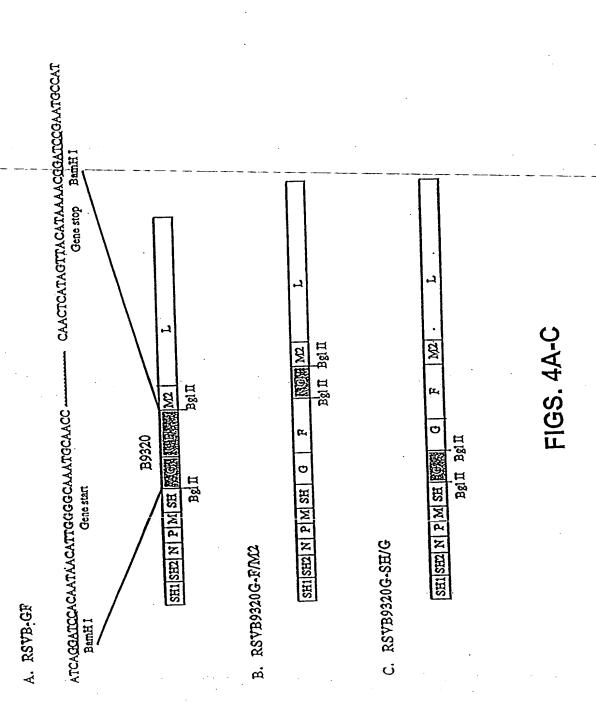


FIG.





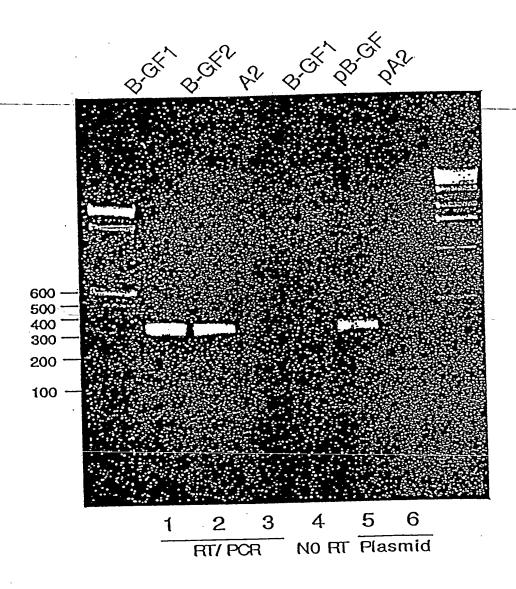
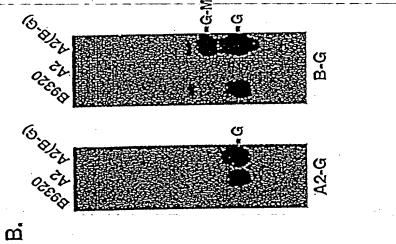
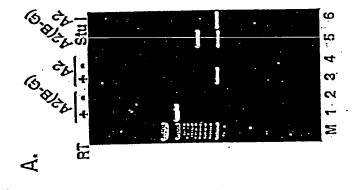


FIG. 5





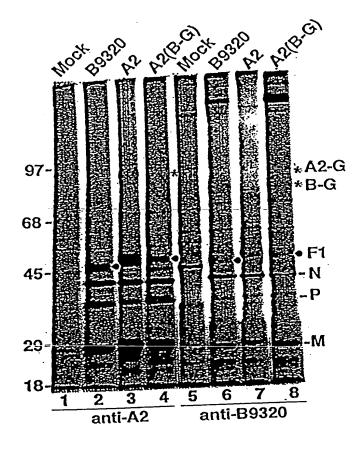


FIG. 7

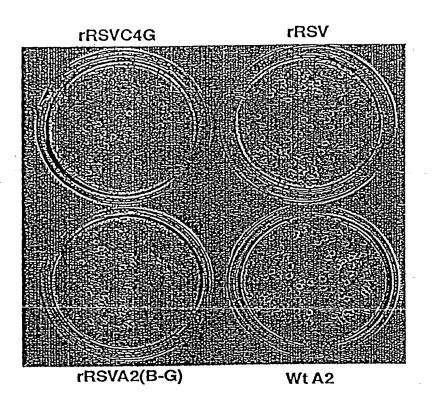


FIG. 8

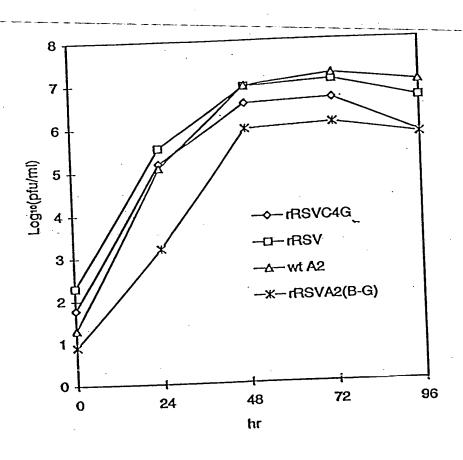


FIG. 9

(Sheet 10 of 28)

Charged. Clusters (Amino Acids that are underlined were changed to alanines) Mutations in cpts-248/404

EEE

TWISNCINTIMKSIG IMNITDAANKAQKNI ERQAMDAVKINGNET

ITEEDQFRKRFYNSM

ELYFLFRIFGHPMVD YXKLNTY PSLLELTE

NIINGRWIILLSKFL

IYRIIKGFVNNYNRW AISPPKNLIWTSFPR LIGKERELSVGRMFA YISKCSIITDL,SKFN

YGDCILKLFHNEGFY

LRCGENNVILTQLFL

LSRVCHTLLDKTVSD YFYLLSSLSMLRGAF VVNOSYLANIPNHVVS SISNKSNRYNDNYAN PYIGDHIVDLANIVDE

RLPKKVDLEMIINDK

TWKDISLSRLNVCLI

KELKKITVITYNQFL IIKEVEGFIMSLILN KLIKLAGDNWLANLS PTLRNAIVLPLRWLT

YTKLNNILTQYRSNE

KDKITKSNNGQDEDNS

DVKVYAILMKLGLKE

LGSYIFNGPYLKNDY TNLLKKIIRRAIEIS

DSYLKGVISFSECNA LMTYKSMTSSEQIAT

MPIINGNSANVYLT

VITTIIKDDILSAVK VRNEGFTLIDNQTLS

KGEIKLEEPTYFQSL

300 375 375 525 600 675 825 900

> RDLIVLSGLRFYREF YLKDNKFNECDLYNC

LKFSESDKSRRVLEY MIAENILQFFPESLT

NYMPSHIQNYIEHEK

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QAFRYETSCICSDVL

ISLKGKFSITALING KTIQHNGVYYPASIK

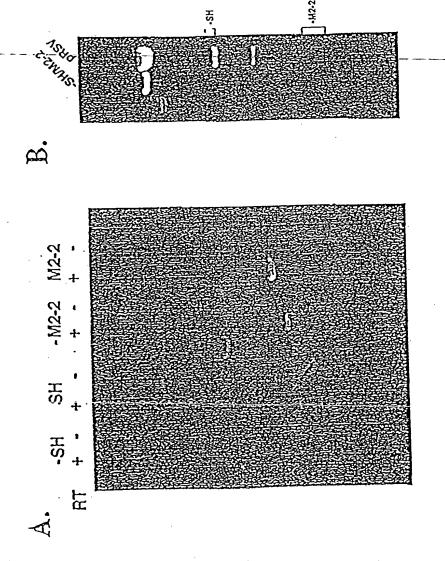
(Sheet 11 of 28)

		•	,	e residues	C Cysteine
2165	NEHNE				HKHIMITKWENHVIN
2100	STAGRNEVFSNKLIN	LSKLKSVVSGDILSY		1 1-4	AKLILSRIKNFIMPK
2025	IGPANIFPVFNVVQN	LGSKLKGSEVYLVLT	DEKLDNITILKTYVC	-	FWSK HVRKCKYCSSV
1950			LTIPATDATMIHMS		RSLKDCNDHSLPIEF
1875	LRTWVELHPDIRYIY	KISIEXILKDLKIKD PNCIAFIGEGAGNLL	KISIEYILKDLKIKD	-	SLVHNSTSLYCMLPW
1800		INDELESNIKKUTKSS AMIRINYSKODLYNI FPMVVIDRIIDHSGN	AMIRINYSKODLYNL		KKTI NDYCIGKNVDS
1725	ETLENILANPIKSND	SDNTHLLTKHIRIAN SELENNYNKLYHPTP		-	TATTORTHIKNKHKEN
1650	MKAILTYIDLVRMGL	TVCPWVNIDYHPTH			K SMSKTFT, BOKVIKY
1575	SDILCVLELIDSSYW	HKGYGKAKLECDMNT	LKVFFNAYKTYLLCF	KDWGEGYITDHAFIN	TI.TTOI.MKDSKGTEE
1500	FHNTYILSTNIAGHW	HVNSNLILAHKISDY	YVELFLSNKTLKSGS	OKOHMFL PDK ISLTO	THEOVITHKLKOVI
1425	ILIPKLNEIHLMKPP	MSVVEQFINVCPNRI	IDIVFONCISFGLSL		A CITO VROMINYHEDY
1350	YLHRLTVSSRPCEFP	YEKAKKLFPQYLSVN YLHRLIVSSRPQEFP	DEFMEELSIGTLGLT		VAR OVE JEKKORDOID
1275	PWVGSSTQEKKTMPV	YNVNSLTRGERGPTK			T. SKYATER SMST.SN
1200	RDKREILSMENLSIT	KNITLLIRILPLDON ROKREILSMENLSIT	AIDLIDIDRATEMMR	1 1-	VECT. DRYKE BKTVNT.
1125	MONIEPTYPHGLRVV	SAQHYTTTEIDIMDI	BEROAKITSEINRIA VTEVISTAPNKIFSK	, U.	MARTINE DOOR OF COURT
1050	LNKFLTCIITFDKNP	TEAIVHSVFILSYYT NHDLKDKLQDLSDDR	TEATVHSVFILSYYT	12	TINATION TO THE GOOD
975	HLKTFFNLDNIDTAL	VWINNOIPIOINNHA LCMMKLYLDIIKVIK HIKTFFMLDNIDIAL	VWLYNQIALQLKNHA		TRAVELECT CST. TOBL
900	KVLRVGPWINTILDD	THIN YENDER GOTT KOTETYISRDMOFMS KTIQHNGVYYPASIK KVIRVGPWINTILDD	KGTETYISRDMOFMS		SWIE TYTE CERTAINS

Cysteine residues that were changed to valine or aspartic acid

Cysteine residue deleted

0 44



FIGS, 12A-B



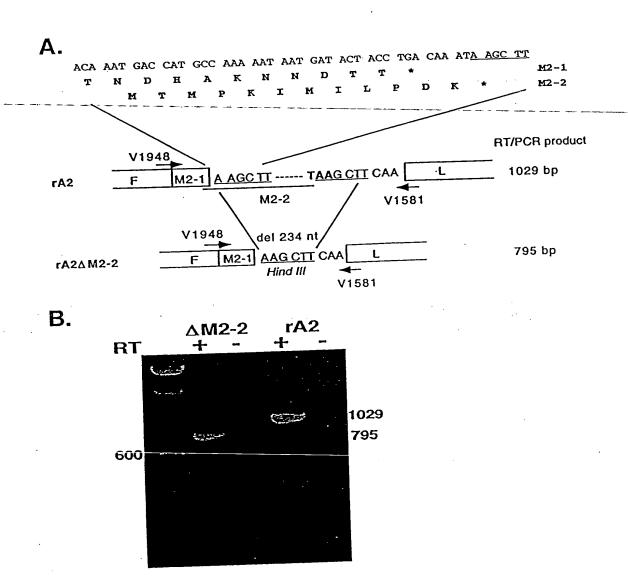


FIG. 13

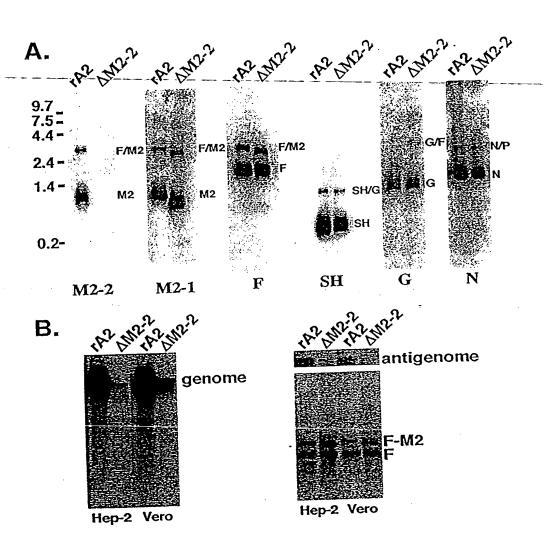


FIG. 14

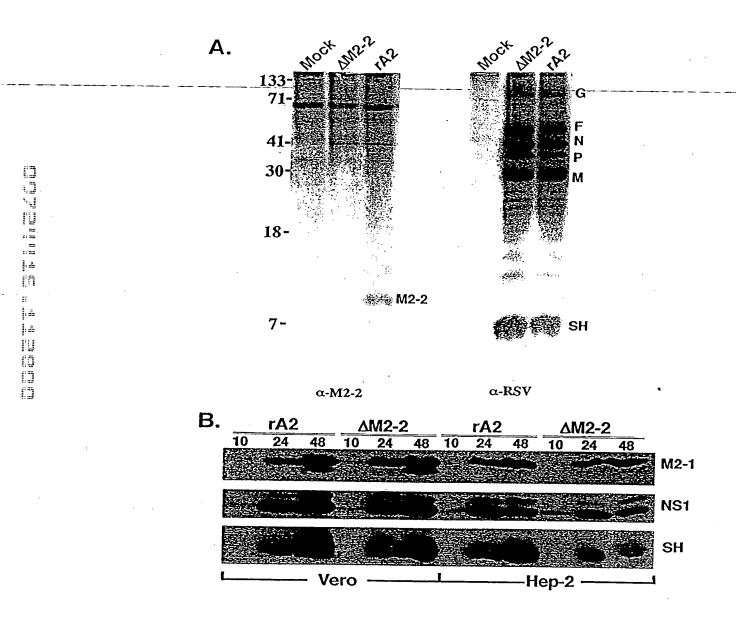


FIG. 15

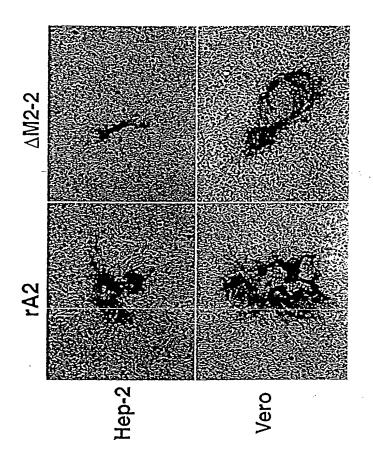
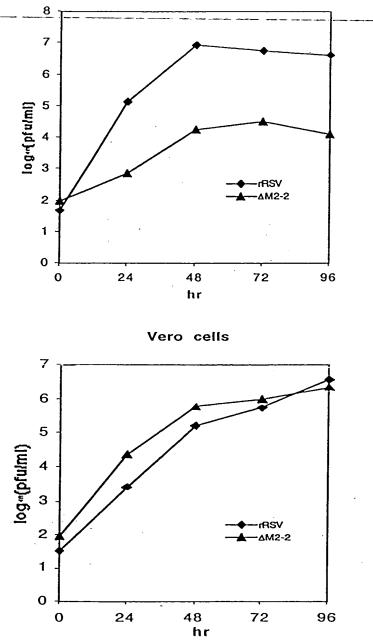


FIG. 16.



Hep-2 cells

FIG. 17

96

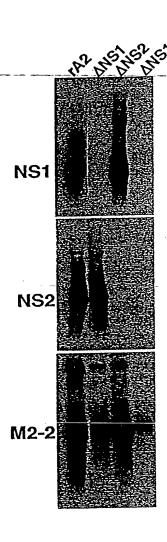


FIG. 18

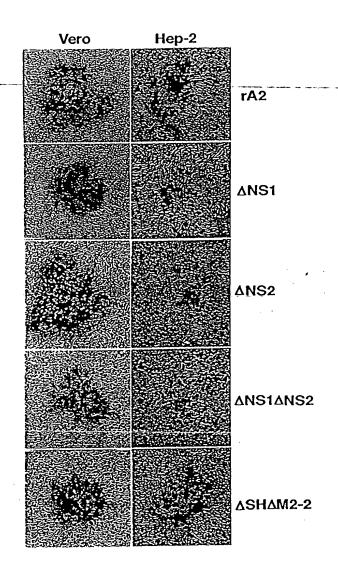


FIG. 19

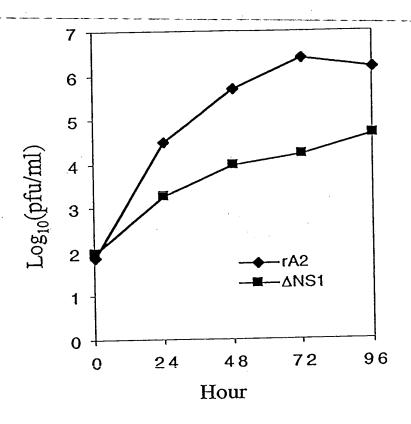


FIG. 20

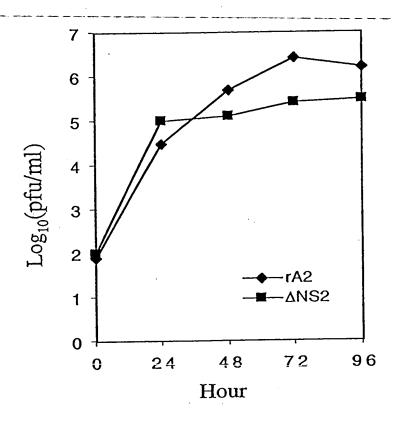


FIG. 21



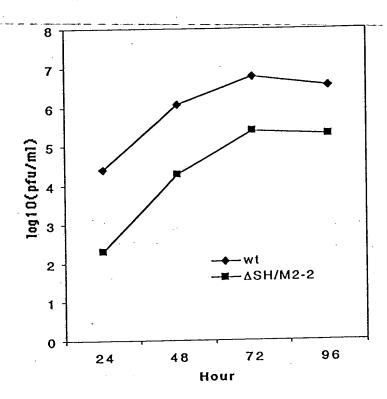


FIG. 22

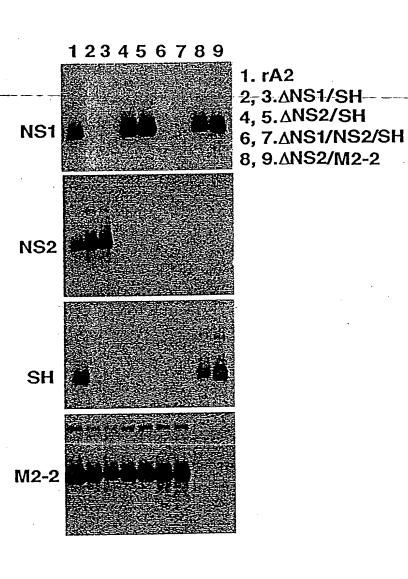


FIG. 23

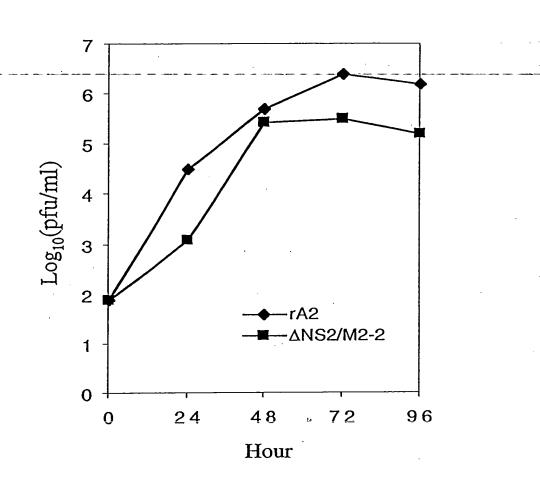


FIG. 24



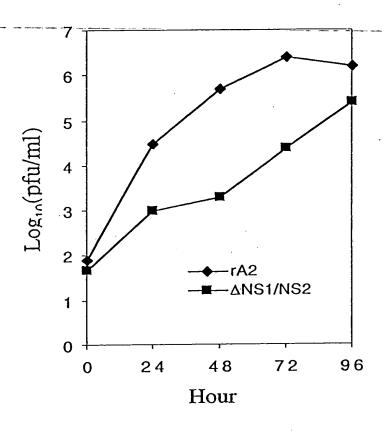
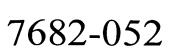


FIG. 25

FIG. 20



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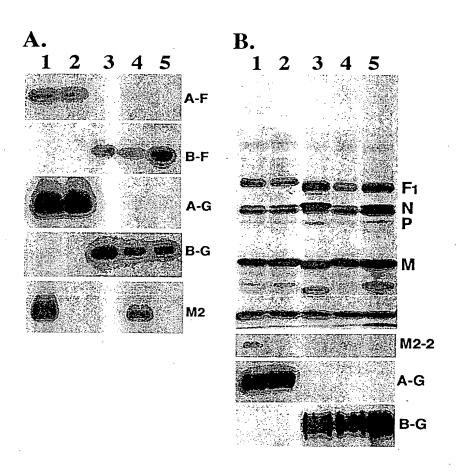


FIG. 27

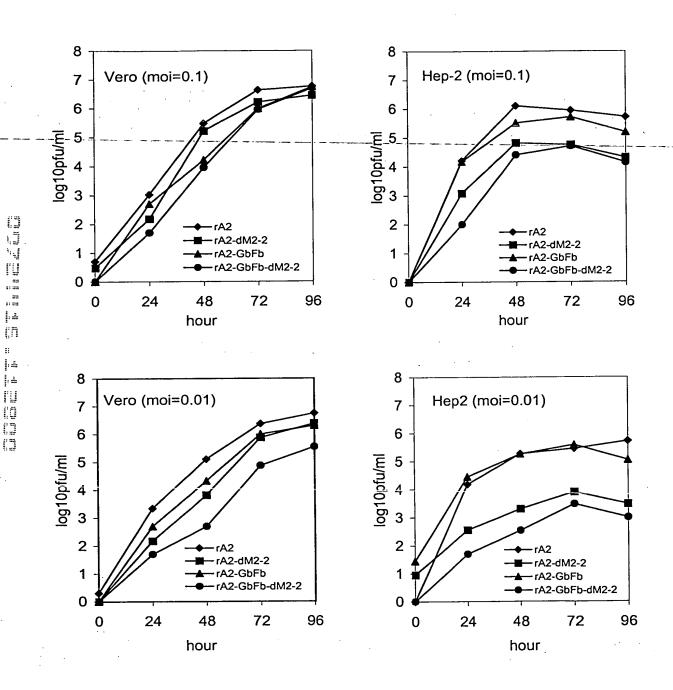


FIG. 28